

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449A/PTO				<b><i>Complete if Known</i></b>	
<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>				Application Number	10/595,485-Conf. #1304
				Filing Date	September 6, 2007
				First Named Inventor	Ezio GHIGO
				Art Unit	1646
				Examiner Name	Not Yet Assigned
Sheet	1	of	6	Attorney Docket Number	

U.S. PATENT DOCUMENTS

## FOREIGN PATENT DOCUMENTS

Examiner Signature		Date Considered	
-----------------------	--	--------------------	--

**\*EXAMINER:** Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. <sup>1</sup> Applicant's unique citation designation number (optional). <sup>2</sup> See Kinda Codes of USPTO Patent Documents at [www.uspto.gov](http://www.uspto.gov) or MPEP 901.04. <sup>3</sup> Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup> For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup> Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. <sup>6</sup> Applicant is to place a check mark here if English language Translation is attached.

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449/PTO				<b>Complete if Known</b>	
				Application Number	<b>10/595,485-Conf. #1304</b>
				Filing Date	September 6, 2007
				First Named Inventor	Ezio GHIGO
				Art Unit	1646
				Examiner Name	Not Yet Assigned
Sheet	2	of	6	Attorney Docket Number	0290494.00122US1

<b>NON PATENT LITERATURE DOCUMENTS</b>				
Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.		
	CA	ARIYASU, H. et al., "Stomach is a Major Source of Circulating Ghrelin, and Feeding State Determines Plasma Ghrelin-Like Immunoreactivity Levels in Humans", The Journal of Clinical Endocrinology & Metabolism, Vol. 86, No. 10 (2001) pp. 4753-4758		T <sup>2</sup>
	CB	BEDNAREK, M. A. et al., "Structure - Function Studies on the New Growth Hormone-Releasing Peptide Ghrelin: Minimal Sequence of Ghrelin Necessary for Activation of Growth Hormone Secretagogue Receptor 1a", J. Med. Chem, Vol. 43 (2000) pp. 4370-4376		
	CC	CAIXAS, A. et al., "Insulin, unlike Food Intake, does not Suppress Ghrelin in Human Subjects", The Journal of Clinical Endocrinology & Metabolism, Vol. 87, No. 4 (2002) pp. 1902-1906		
	CD	CHAPMAN, I. et al. "Enhancement of Pulsatile Growth Hormone Secretion by Continuous Infusion of a Growth Hormone-Releasing Peptide Mimetic, L-692,429, in Older Adults - A Clinical Research Center Study." Journal of Clinical Endocrinology and Metabolism, Vol. 81, No. 8 (1996) pp. 2874-2880		
	CE	CHAPMAN, I. et al., "Oral Administration of Growth Hormone (GH) Releasing Peptide-Mimetic MK-677 Stimulates the GH/Insulin-Like Growth Factor-I Axis in Selected GH-Deficient Adults", Journal of Clinical Endocrinology and Metabolism, Vol. 82, No. 10 (1997) pp. 3455-3463		
	CF	CUMMINGS, D. E. et al., "A Preprandial Rise in Plasma Ghrelin Levels Suggests a Role in Meal Initiation in Humans", Diabetes, Vol. 50 (August 2001) pp. 1714-1719		
	CG	DATE, Y. et a., "Ghrelin Acts in the Central Nervous System to Stimulate Gastric Acid Secretion", Biochemical and Biophysical Research Communications, Vol. 280 (2001) pp. 904-907		
	CH	DATE, Y. et al., "Ghrelin is Present in Pancreatic $\alpha$ -Cells of Humans and Rats and Stimulates Insulin Secretion", Diabetes, Vol. 51 (January 2002) pp. 124-129		
	CI	DATE, Y. et al., "Ghrelin, a Novel Growth Hormone-Released Acylated Peptide, Is Synthesized in a Distinct Endocrine Cell Type in the Gastrointestinal Tracts of Rats and Humans." Endocrinology, Vol. 141, No. 11 (2000) pp. 4255-4261		
	CJ	European Office Action, corresponding to European Application No. 02787266.2, October 7, 2008		

Examiner Signature	Date Considered
--------------------	-----------------

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>1</sup>Applicant's unique citation designation number (optional). <sup>2</sup>Applicant is to place a check mark here if English language Translation is attached.

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449/PTO				<b>Complete if Known</b>	
				Application Number	<b>10/595,485-Conf. #1304</b>
				Filing Date	September 6, 2007
				First Named Inventor	Ezio GHIGO
				Art Unit	1646
				Examiner Name	Not Yet Assigned
Sheet	3	of	6	Attorney Docket Number	0290494.00122US1

<b>NON PATENT LITERATURE DOCUMENTS</b>				
Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.		T <sup>2</sup>
	CK	FURUSE, M. et al., "Intracerebroventricular injection of ghrelin and growth hormone releasing factor inhibits food intake in neonatal chicks", Neuroscience Letters, Vo. 301 (2001) pp. 123-126		
	CL	GUALILLO et al. "Ghrelin, A Novel Placental-Derived Hormone", Endocrinology, Vol. 142, No. 2, (2001) pp. 788-794		
	CM	HATTORI et al. "GH, GH Receptor, GH Secretagogue Receptor, and Ghrelin Expression in Human T Cells, B Cells, and Neutrophils." The Journal of Clinical Endocrinology and Metabolism, Vol. 86, No. 9 (2001) pp. 4284-4291		
	CN	HORVATH, T. L. et al., "Minireview: Ghrelin and the Regulation of Energy Balance - A hypothalamic Perspective", Endocrinology, Vol. 142, No. 10 (2001) pp. 4163-4169		
	CO	INUI, Akio, "Ghrelin: An orexigenic and Somatotropic Signal from the Stomach", Nature Reviews, Vol. 2 (August 2001) pp. 1-10		
	CP	KAMEGAI, J. et al., "Central Effect of Ghrelin, an Endogenous Growth Hormone Secretagogue, on Hypothalamic Peptide Gene Expression", Endocrinology, Vol. 141, No. 12, 2000, pp. 4797-4800		
	CQ	KAMEGAI, J. et al., "Chronic Central Infusion of Ghrelin Increases Hypothalamic Neuropeptide Y and Agouti-Related Protein mRNA Levels and Body Weight in Rats", Diabetes, Vol. 40 (November 2001) pp. 2438-2443		
	CR	KOJIMA et al. "Ghrelin is a growth-hormone-releasing acylated peptide from stomach" Nature, Vol. 402, 9 December 1999. (pp 656-660)		
	CS	KORBONITS et al. "The presence of ghrelin in normal and adenomatous human pituitary," Department of Endocrinology, St. Bartholomew's Hospital, London, UK and Department of Biochemistry, National Cardiovascular Center Institute, Fujishirodau, Suita, Osaka Japan (11 pgs)		
	CT	LUCIDI, P. et al., "Ghrelin is not Necessary for Adequate Hormonal Counterregulation of Insulin-Induced Hypoglycemia", Diabetes, Vol. 51 (October 2002) pp. 2911-2914		

Examiner Signature	Date Considered
--------------------	-----------------

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>1</sup>Applicant's unique citation designation number (optional). <sup>2</sup>Applicant is to place a check mark here if English language Translation is attached.

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449/PTO				<b>Complete if Known</b>	
				Application Number	<b>10/595,485-Conf. #1304</b>
				Filing Date	September 6, 2007
				First Named Inventor	Ezio GHIGO
				Art Unit	1646
				Examiner Name	Not Yet Assigned
Sheet	4	of	6	Attorney Docket Number	0290494.00122US1

<b>NON PATENT LITERATURE DOCUMENTS</b>			
Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>2</sup>
	CU	MASUDA, Y. et al., "Ghrelin Stimulates Gastric Acid Secretion and Motility in Rats", Biochemical and Biophysical Research Communications, Vol. 276 (2000) pp. 905-908	
	CV	MATSUMOTO, M. et al., "Structure-Activity Relationship of Ghrelin: Pharmacological Study of Ghrelin Peptides", Biochemical and Biophysical Research Communications, Vol. 287 (2001), pp. 142-146	
	CW	MORI, K. et al., "Kidney produces a novel acylated peptide, ghrelin", FEBS Letters, Vol. 486 (2000) pp. 213-216	
	CX	MUCCIOLO, G. et al. "Neuroendocrine and peripheral activities of ghrelin: implications in metabolism and obesity," European Journal of Pharmacology, Vol. 440 (2002) pp. 235-254	
	CY	MUCCIOLO, G. et al., "Binding of 125I-labeled ghrelin to membranes from human hypothalamus and pituitary gland", J. Endocrinol. Invest., Vol. 24 (2001) pp. RC7-RC9	
	CZ	MULLER, A. F. et al., "Blockade of the Growth Hormone (GH) Receptor Unmasks Rapid GH-Releasing Peptide-6-Mediated Tissue-Specific Insulin Resistance", The Journal of Clinical Endocrinology & Metabolism, Vol. 86, No. 2, 2001, pp. 590-593	
	CA1	NAGAYA, N. et al., "Hemodynamic and hormonal effects of human ghrelin in healthy volunteers", Am J. Physiol Regulatory Integrative Comp. Physiol., Vol. 280 (2001) pp. R1483-R1487	
	CB1	NAKAGAWA, E. et al., "Hyperglycaemia suppresses the secretion of ghrelin, a novel growth-hormone-releasing peptide: responses to the intravenous and oral administration of glucose", Clinical Science, Vol. 103 (2002) pp. 325-328	
	CC1	OKUMURA, H. et al., "Vasodilatory Effect of Ghrelin, an Endogenous Peptide from the Stomach", Journal of Cardiovascular Pharmacology, Vol. 39 (2002) pp. 779-783	
	CD1	PAGOTTO, U. et al., "Plasma Ghrelin, Obesity, and the Polycystic Ovary Syndrome: Correlation with Insulin Resistance and Androgen Levels", The Journal of Clinical Endocrinology & Metabolism, Vol. 87, No. 12 (2002) pp. 5625-5629	

Examiner Signature	Date Considered
--------------------	-----------------

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>1</sup>Applicant's unique citation designation number (optional). <sup>2</sup>Applicant is to place a check mark here if English language Translation is attached.

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449/PTO				<b>Complete if Known</b>	
				Application Number	<b>10/595,485-Conf. #1304</b>
				Filing Date	September 6, 2007
				First Named Inventor	Ezio GHIGO
				Art Unit	1646
				Examiner Name	Not Yet Assigned
Sheet	5	of	6	Attorney Docket Number	0290494.00122US1

<b>NON PATENT LITERATURE DOCUMENTS</b>				
Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.		T <sup>2</sup>
	CE1	SAAD, M. et al., "Insulin Regulates Plasma Ghrelin Concentration", The Journal of Clinical Endocrinology & Metabolism, Vol. 87, No. 8 (2002) pp. 3997-4000		
	CF1	SVENSSON, J. et al., "Two-Month Treatment of Obese Subjects with the Oral Growth Hormone (GH) Secretagogue MK-677 Increases GH Secretion, Fat-Free Mass, and Energy Expenditure", Journal of Clinical Endocrinology and Metabolism. Vol. 83, No. 2 (1998) pp. 362-369		
	CG1	TANKA, M. et al., "Testis-specific and developmentally induced expression of a ghrelin gene-deprived transcript that encodes a novel polypeptide in the mouse", Biochimica et Biophysica Acta, Vol. 1522 (2001) pp. 62-65		
	CH1	TENA-SEMPERE, M. et al., "Novel Expression and Functional Role of Ghrelin in Rat Testis", Endocrinology, Vol. 143, No. 2, 2002, pp. 717-725		
	CI1	TOLLE, V. et al., "Ultradian Rhythmicity of Ghrelin Secretion in Relation with GH, Feeding Behavior, and Sleep-Wake Patterns in Rats", Endocrinology, Vol. 143, No. 4 (2002) pp. 1353-1361		
	CJ1	TSCHOP, M. et al., "Ghrelin induces adiposity in rodents", Letters to Nature, Vol. 407 (19 October 2000) pp. 908-913		
	CK1	VOLANTE, et al. "Extensive DNA Fragmentation in Oxyphilic Cell Lesions of the Thyroid," The Journal of Histochemistry and Cytochemistry, Vol. 49, No. 8 (2001) pp. 1003-1011		
	CL1	VOLANTE, M. et al., "Expression of Ghrelin and of the GH Secretagogue Receptor by Pancreatic Islet Cells and Related Endocrine Tumors", The Journal of Clinical Endocrinology & Metabolism, Vol. 87, No. 3 (2002) pp. 1300-1308		
	CM1	VOLANTE, M. et al., "Ghrelin Expression in Fetal, Infant, and Adult Human Lung", The Journal of Histochemistry & Cytochemistry, Vol. 50, No. 8 (2002) pp. 1013-1021		
	CN1	WIERUP, N. et al., "The ghrelin cell: a novel developmentally regulated islet cell in the human pancreas", Regulatory Peptides, Vol. 107 (2002) pp. 63-69		

Examiner Signature	Date Considered
--------------------	-----------------

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>1</sup>Applicant's unique citation designation number (optional). <sup>2</sup>Applicant is to place a check mark here if English language Translation is attached.

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449/PTO				<b><i>Complete if Known</i></b>	
				Application Number	<b>10/595,485-Conf. #1304</b>
				Filing Date	September 6, 2007
				First Named Inventor	Ezio GHIGO
				Art Unit	1646
				Examiner Name	Not Yet Assigned
Sheet	6	of	6	Attorney Docket Number	0290494.00122US1

<b>NON PATENT LITERATURE DOCUMENTS</b>			
Examiner Initials	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>2</sup>
	CO1	WREN, A. M. et al., "Ghrelin Enhances Appetite and Increases Food Intake in Humans", The Journal of Clinical Endocrinology & Metabolism, Vol. 86, No. 12 (2001) pp. 5992-5995	
	CP1	WREN, A.M. et al., "The Novel Hypothalamic Peptide Ghrelin Stimulates Food Intake and Growth Hormone Secretion", Endocrinology, Vol. 141, No. 11 (2000) pp. 4325-4328	
	CQ1	YOSHIHARA, F. et al., "Ghrelin: a novel peptide for growth hormone release and feeding regulation", Genes and Nutrition, pp. 391-395	
	CR1	ZHANG, W. et al., "Inhibition of pancreatic protein secretion by ghrelin in the rat", Journal of Physiology, Vol. 537, No. 1 (2001) pp. 231-236	

Examiner Signature	Date Considered
--------------------	-----------------

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>1</sup>Applicant's unique citation designation number (optional). <sup>2</sup>Applicant is to place a check mark here if English language Translation is attached.